

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In application of:)	
	Ole Thastrup et al.)	
Serial No.	10/072,036)	Art Unit
)	1633
Confirmation No.	3012)	
For:	A METHOD FOR EXTRACTING QUANTITATIVE INFORMATION RELATING TO AN INFLUENCE ON A CELLULAR RESPONSE)	
Filing Date:	February 5, 2002)	
Examiner:	M.D. Burkhardt)	
Customer No.	022913)	

DECLARATION OF CHRIS M. IRELAND. PH.D.**UNDER 37 C.F.R. & 1.132**

Mail Stop AMENDMENT
Commissioner for Patents P.O.
Box 1450
Alexandria, VA 22313-1450

Dear Sir:

I, Chris M. Ireland, Ph.D., hereby declare as follows:

1. I am personally knowledgeable of the facts stated herein.
2. I am a professor and chairman of the Department of Medicinal Chemistry in the College of Pharmacy at the University of Utah, Salt Lake City, Utah. I have significant experience in preparing compounds and screening compounds for biological activity (*see* Appendix A: Curriculum Vitae of Chris M. Ireland). Additionally, I am knowledgeable in the field of developing and validating high throughput screening (HTS) assays for biological activity and drug development, and also knowledgeable in the field of formatting and processing of chemical diversity libraries for use in HTS assays. I am therefore knowledgeable in the field preparing and screening compounds for biological activity, such as disclosed and claimed in the U.S. Patent Application Serial No.10/072,036 ("Subject Application"), which is currently under examination.

3. I have reviewed and understand the Subject Application and the *Carey* reference cited by the Examiner in the Office Action of October 20, 2006.

4. I am not an inventor of the subject matter disclosed and claimed in the Subject Application.

5. I am not currently employed by and have never been employed or associated with Fisher BioImage ApS or Thermo Fisher Scientific.

6. I do not have a personal interest in the Subject Application.

7. A dictionary definition of "library" reads as follows: "[a] place in which literary and artistic materials, such as books, periodicals, newspapers, pamphlets, prints, records, and tapes, are kept for reading, reference, or lending"; "[a] collection of such materials, especially when systematically arranged"; "[a] room in a private home for such a collection"; "[a]n institution or foundation maintaining such a collection"; "[a] commercial establishment that lends books for a fee"; "[a] series or set of books issued by a publisher"; "[a] collection of recorded data or tapes arranged for ease of use"; "[a] set of things similar to a library in appearance, function, or organization: *a library of computer programs*"; and "*Genetics* A collection of cloned DNA sequences whose location and identity can be established by mapping the genome of a particular organism." (See, *The American Heritage® Dictionary of the English Language, Fourth Edition*. Houghton Mifflin Company, 2004. 21 Feb. 2007. <Dictionary.com <http://dictionary.reference.com/browse/library>>.)

8. A dictionary definition of "compounds" with respect to the field of chemistry reads as follows: "[a] pure, macroscopically homogeneous substance consisting of atoms or ions of two or more different elements in definite proportions that cannot be separated by physical means. A compound usually has properties unlike those of its constituent elements." (See, *The American Heritage® Dictionary of the English Language, Fourth Edition*. Houghton Mifflin Company, 2004. 23 Feb. 2007. <Dictionary.com <http://dictionary.reference.com/browse/compound>>.)

9. The ordinary meaning of a "library of compounds," as evidenced by the aforementioned dictionary definitions, is a collection of compounds that are either pure or present at a known concentration, and such compounds of the library of compounds are arranged for ease of use. Additionally, the plain meaning of a "library of compounds" is similar to a standard book library in appearance, function, or organization.

10. As used in the Subject Application, a "library of compounds" is a term of art that is well known to those of ordinary skill in the art of screening compounds. A skilled artisan in the art of screening compounds would understand a "library of compounds" to be a collection of compounds that

are either pure or present at a known concentration, and arranged so that each compound can be selected from the collection of compounds for use in an experiment either alone or in combination with other compounds of the library. Also, a skilled artisan would understand that a library of compounds is a collection of compounds assembled for the purpose of testing the compounds to determine suitability for a particular purpose, or to detect wanted or unwanted attributes of the compounds. Additionally, a skilled artisan would understand that a "library of compounds" has a function similar to a standard book library in that each compound of the collection of compounds is individually retained within a container such that a single compound of the collection of compounds can be individually selected for use similar to how a single book can be selected in a standard book library. The compounds can be individually retained in the container in a pure state or in a specific, known concentration in a specific solvent that is compatible with the environment in which the compound will be used, wherein the container is substantially devoid of any contaminating compounds or substances. Furthermore, a skilled artisan would understand that more than one compound of a "library of compounds" can be selected as a group for use similarly to how a group of books can be selected in a standard book library.

11. A dictionary definition of "screening" reads as follows: "[t]o examine (a job applicant, for example) systematically in order to determine suitability"; "[t]o test or evaluate (a student) to determine placement in an educational system or to identify specific learning needs"; "[t]o test or examine for the presence of disease or infection: *screen blood; screen a patient*"; "[t]o subject to genetic screening"; and "[a] systematic examination or assessment, done especially to detect an unwanted substance or attribute. (See, *The American Heritage® Dictionary of the English Language, Fourth Edition*. Houghton Mifflin Company, 2004. 23 Feb. 2007. <Dictionary. com <http://dictionary.reference.com/browse/screening>>.)

12. The ordinary meaning of "screening a library of compounds," as evidenced by the aforementioned dictionary definitions, is a process to systematically examine, test, or evaluate the compounds or a combination of compounds of a library of compounds to determine suitability for a particular purpose, or detect wanted or unwanted attributes of the compound.

13. As used in the Subject Application, "screening a library of compounds" is a term of art for an experimental process that is well known to those of ordinary skill in the art of screening compounds. A skilled artisan in the art of screening compounds would understand "screening a library of compounds" to be a process to systematically examine, test, or evaluate the compounds of the library of compounds in order to determine suitability for a particular purpose, or to detect wanted or

unwanted attributes of the compounds. Additionally, a skilled artisan in the art of screening compounds would understand that "screening a library of compounds" would include a process to systematically examine, test, or evaluate whether or not a known concentration of a compound or a combination of compounds of the library of compounds is biologically active, and so that the biological activity can be correctly attributed to a known concentration of the compound. Furthermore, such "screening a library of compounds" inherently includes comparing a compound or a combination of compounds of the library of compounds to a control so that the presence or absence of any biological activity is correctly attributed to the compound or combination of compounds.

14. The specification of the Subject Application (PG-PUB 2003/0082564) discloses a "library of compounds" at the following paragraphs: [0027] and [0103]. Additionally, the specification of the Subject Application (PG-PUB 2003/0082564) discloses a "screening" at paragraphs [0001], [0013], [0027], [0036], [0058-0063], and [0126-0141]. The Examples and Figures describe protocols and illustrate results for such a screening. Thus, the Subject Application discloses "screening a library of components."

15. The *Carey* reference does not disclose a "library of compounds" or a group of compounds that could be construed as a "library of compounds" because there is no reference to a collection of compounds that are each either pure or present at a known concentration, and that are assembled for the purpose of testing the compounds to determine suitability for a particular purpose, or to detect wanted or unwanted attributes of the compounds.

16. A skilled artisan would not consider "any and all substances [or compounds] mentioned in the Materials and Methods section on pages 986-987" in the *Carey* reference to be a library of compounds. In part, this is because the substances of *Carey* are not described to have each compound of the substance to be individually retained within a container at a known concentration such that a single compound of the substance can be individually selected. Additionally, the Materials and Methods section of *Carey* does not teach or suggest the compounds recited therein are a "library of compounds" that are assembled for the purpose of testing the compounds to determine suitability for a particular purpose, or to detect wanted or unwanted attributes of the compounds.

17. A skilled artisan would not consider dexamethasone, phenol red, serum, and charcoal-stripped serum to be a library of compounds because only dexamethasone could be construed to be a compound that is either pure or present at a known concentration. Also, *Carey* does not teach or suggest that phenol red is available for being selected in the absence of cell culture media, and thereby phenol red is not taught or suggested to be present in a pure state or in a specific, known concentration

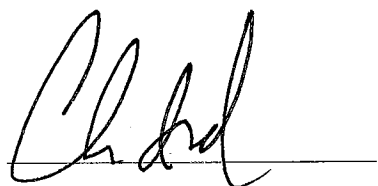
in a specific solvent that is compatible with the environment in which the compound will be used, wherein the solvent is substantially devoid of any contaminating compounds or substances because the other compounds contained in media would be considered to be contaminants. Additionally, neither serum nor charcoal-stripped serum is a pure compound, and the compounds contained therein are not all present at known concentrations.

18. A skilled artisan would not consider the compounds contained in serum or charcoal-stripped serum to be a library of compounds because the compounds contained therein are not pure and are not all present at known concentrations because serum contains many compounds at unknown concentrations. Also, the compounds of serum are not a collection of compounds that are individually retained within a container such that a single compound can be individually selected.

19. The *Carey* reference does not disclose a method of "screening a library of compounds," or reference any experiment (*e.g.*, experiments conducted with dexamethasone, phenol red, serum, and/or charcoal-stripped serum) that a skilled artisan would construe as "screening a library of compounds" because a "library of compounds" is not taught or suggested as discussed above. Additionally, only dexamethasone could be construed to be in a pure state or at a known concentration that is included in a process to be systematically tested to determine suitability for a particular purpose, or to detect wanted or unwanted attributes, and testing dexamethasone alone does not constitute "screening a library of compounds."

20. I declare further that all statements made herein of my own knowledge are true and that all statements are made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful, false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code, and that such willful, false statements may jeopardize the validity of the application or any patent issuing thereon.

Signed this 20th day of March, 2007.


CHRIS M. IRELAND, Ph.D

APPENDIX A

CURRICULUM VITAE

CHRIS M. IRELAND

**University of Utah
College of Pharmacy
Department of Medicinal Chemistry
Salt Lake City, Utah 84112
(801) 581-8305**

Birth date

November 22, 1951, San Diego, California, U.S. Citizen

Marital Status

Married, four children

Education

1973	B.A. University of California, San Diego
1977	Ph.D. Scripps Institution of Oceanography

Position

Professor and Chair, Department of Medicinal Chemistry

Professional/Research Experience

2002-	Chair, Department of Medicinal Chemistry
1991-	Adjunct Professor, Department of Chemistry, University of Utah
1991-	Professor, Department of Medicinal Chemistry, College of Pharmacy, University of Utah
1986-91	Adjunct Associate Professor, Department of Chemistry, University of Utah
1986-91	Associate Professor, Department of Medicinal Chemistry, College of Pharmacy, University of Utah
1983-86	Adjunct Assistant Professor, Department of Chemistry, University of Utah

1983-86	Assistant Professor, Department of Medicinal Chemistry, College of Pharmacy, University of Utah
1980-83	Assistant Professor, University of Connecticut
1980	Postdoctoral Fellow, University of Illinois
1978-80	NIH Postdoctoral Fellow, University of Hawaii

Professional/Research Interests

Marine pharmacognosy, natural products chemistry, structure and mechanism of action of antitumor natural products, marine microbiology, structure and neurotoxicity of conotoxins

Scientific/Professional Societies

American Chemical Society-Organic Division
 American Society of Pharmacognosy
 American Association of Cancer Researchers

Honors and Distinctions

2007	University of Utah, Distinguished Scholarly & Creative Research Award
2006	University of Utah, Distinguished Graduate Research Mentor Award
2006	College of Pharmacy, University of Utah, Distinguished Teaching Award
1987-92	National Institutes of Health Research Career Development Award
1985-89	Alfred P. Sloan Foundation Fellow
1981	University of Connecticut Summer Faculty Fellow
1978-80	National Institutes of Health Postdoctoral Fellow

National Public Service Activities and Committee Memberships

American Society of Pharmacognosy, Chairman, Awards and Funds Committee, 1985-88
 American Society of Pharmacognosy, Chairman & Host, 1988 Annual Meeting
 Member Bio-organic and Natural Products study section-NIH, 1989-1993
 Vice-President/President Elect, American Society of Pharmacognosy, 1990-91
 Co-Chairman, Marine Natural Products Gordon Conference, 1990
 President, American Society of Pharmacognosy, 1991-1992
 Chairman, Marine Natural Products Gordon Conference, 1992
 American Society of Pharmacognosy, Chair, Scientific Committee & Member, Local Committee, 1993 Annual Meeting
 American Society of Pharmacognosy, Chair, Personnel Committee, 1993-1996

NIH DRG Reviewers Reserve, 1993- Have served ad hoc on numerous study sections and site visit teams.

American Society of Pharmacognosy, Chair, Research Achievement Award Committee, 1998.

American Society of Pharmacognosy, Chair, Honorary Membership Committee, 2001-2004.

Journal Review Activities:

Tetrahedron Letters
Journal of the American Chemical Society
Journal of Organic Chemistry
Journal of Natural Products
Tetrahedron
Journal of Medicinal Chemistry
Marine Drugs
Organic Letters

Editorial Board Membership

Journal of Natural Products	1999-present
Philippine Journal of Science	1999-present
Marine Drugs	2002-present

Consultant/Advisory Positions (*past* and present)

Member, Scientific Advisory Board, Mycosynthetix, Inc.
Member, Scientific Advisory Board, INI Health
Member, Scientific Advisory Board, OSI Pharmaceuticals
Member, Scientific Advisory Board, NPS Pharmaceuticals
Member, Scientific Advisory Board, Ancile Pharmaceuticals

University Service Activities and Committee Memberships

Combined Programs in Biological Chemistry and Molecular Biology, Executive Committee (University)
Incentive Seed Grants Program (University)
Curriculum Committee (College)
Computer Advisory Committee (College)
Instrument Room Committee (College)
Scholastic Standards Committee (College)
Awards and Funds Committee (College)
Chair of Search Committee for Chairperson of the Department of Pharmacology and Toxicology (College)
NMR Committee (Department)

Department Teaching Committee (Department)
Biological Chemistry Program steering committee (University)
College of Pharmacy Executive Committee (College)

University Teaching/Advisory Committees

Courses taught/new courses developed

PHARM SCI 410	Basics in Pharmaceutical Sciences
PHARM SCI 411	Basics in Pharmaceutical Sciences
PHARM SCI 5113	Basics in Pharmaceutical Sciences
MED CHEM 556	Physiological Chemistry
MED CHEM 561	Natural Products
MED CHEM 601	Instrumental Methods
MED CHEM 789	Seminars
MED CHEM 652	Natural Product(Grad)

Research Faculty Supervised

Rohan Davis, Ph.D., Research Assistant Professor
Timothy S. Bugni, Ph.D., Research Assistant Professor

Post Doctoral Students Supervised-Current Position

Deborah M. Roll, Ph.D.-Senior Scientist, Wyeth Ayerst Research
Tadeusz F. Molinski, Ph.D.-Professor, Department of Chemistry, UC-San Diego
Bradley S. Davidson, Ph.D.-Associate Professor, Department of Chemistry, Utah State University
Robert R. West, Ph.D.-Senior Scientist, Zymogenetics
Brent D. Copp, Ph.D.-Senior Lecturer, Department of Chemistry, University of Auckland
Jodi A. Laakso, Ph.D.-Research Scientist, New Chemical Entities, Inc.
Tommaso A. Foderaro, Ph.D.-Research Scientist, Ciba Gigy
Todd L. Capson, Ph.D.-Research Assistant Professor, Department of Biology, University of Utah
Debra Venables, Ph.D.-Research Scientist, AstraZeneca-Griffith Univeristy
Cameron J. Smith, Ph.D.-Senior Scientist, Merck
Deniz Tasdemir, Ph.D.- Lecturer, University of London
Rohan Davis, Ph.D. - Assistant Lecturer, Griffith University
Damian Laird, Ph.D. - Research Associate, Murdoch University
Camille Chevalier, Ph.D. - Present
Anokha Ratnayake, Ph.D. - Senior Scientist, Wyeth
Daniel LaBarbera, Ph.D. - Present
Kaleem Mohammad, Ph.D. - Present
Rachel Jadulco, Ph.D. - Present

Malcolm McCulloch

Students Graduated

Joseph E. Biskupiak, Ph.D., Associate Professor of Pharmacy Practice, University of Utah
David F. Sestin, Ph.D. - Senior Scientist, Shaman Pharmaceuticals
T. Mark Zabriskie, Ph.D. - Associate Professor, College of Pharmacy, Oregon State University
Tawnya C. McKee, Ph.D. - Staff Research Fellow, National Cancer Institute
Mark P. Foster, Ph.D. - Associate Professor, Department of Biochemistry, Ohio State University
Leonard A. McDonald, Ph.D. - Research Scientist, Wyeth Ayerst Research
J. Christopher Swersey, M.S. - Brewmaster
Annette M. Fernandez, Ph.D. - Bristol Myers Squibb
Scott S. Mitchell, Ph.D. - Senior Scientist, Nereus Pharmaceuticals
Ryan VanWagoner, Ph.D. - Research Assistant Professor, UNC-Wilmington
Robyn James, M.S. - Research Associate, Birnham Insititute
Adam Richardson, Ph.D. - Postdoctoral Fellow, Birnham Insititute
Tim Bugni, Ph.D. - Research Assistant Professor, University of Utah
Sheryl Verbitski, Ph.D. - Senior Scientist, ChromaDex
Imelda Sandoval, Ph.D. - Postdoctoral Fellow, Huntsman Cancer Institute

Current Graduate Students Supervised

Emily Whitson (4rd year)
Charles Veltri (4rd year)

Visiting Scholars Supervised

Imelda Sandoval - University of the Philippines
Giselle Concepcion - University of the Philippines
Teatulohi Matianaho - University of Papua New Guinea
Brent Copp - Auckland University
Manah Dindi - University of Papua New Guinea

Grant Awards History (Current awards are in italics)

NIH Grant 1 S15 CA46245-01; "Silicon Graphics Iris 3130 Work Station," September 30, 1987 to September 29, 1988, \$22,888

College of Pharmacy - BRSG; "Cultured Blue-Green Algae" Sept. 1, 1985 to August 30, 1986, \$3,500

Alfred P. Sloan Foundation; September 16, 1985 to September 16, 1987, \$25,000

University of Utah Research Committee; "Antiviral Agents from Marine Organisms"
November 1, 1983 to October 21, 1984, \$5,000

NIH Grant CA29821; "Antineoplastic Agents from Marine Organisms" May 1, 1982 to
August 31, 1983, \$86,449

PRF Grant 12730-G4; "Photosynthetic Molluscs" September 1, 1980 to August 31, 1982,
\$10,000

UCRF Grant 35-420; "Marine Molluscs" September 1, 1980 to August 31, 1981, \$9,490

NIH Grant CA01179; RCDA April 1, 1987 to March 31, 1992, \$256,000 (direct costs)

NIH Grant CA50750; "National Cooperative Natural Product Drug Discovery Group"
September 1, 1989 to June 30, 1994, Co-Investigator, total budget \$2,500,000

NIH Grant S10 RR06262; "500 MHZ NMR Spectrometer" July 18, 1991 to July 17, 1992
\$400,000 (total costs)

California Sea Grant College Grant S145-N-N; "Technology for Discovery of Anti-
Cancer Drugs from Marine Sources" October 1, 1995 to September 30, 1998, Co-
Investigator, \$195,793 (total costs)

DOC, NOAA Grant NA36RG0036; "Metabolite Studies with Marine Microorganisms"
July 1, 1996 to June 30, 1999, \$310,000.

NIH Grant "Varian INOVA 600 MHZ NMR Spectrometer" April 1, 1999, to March 31,
2000, \$470,732 (total costs).

NIH Grant CA67786; "Anticancer Agents from Unique Natural Products Sources"
September 8, 1995 to August 31, 2000, \$2,348,308 (total costs).

NIH Grant CA67786; "Anticancer Agents from Unique Natural Products Sources"
September 8, 2000 to August 31, 2005, \$4,360,000 (total costs).

Fogarty – FIRCA R03TW06008-01, HIV and TB Screening of PNG Marine Organisms
08/15/2002 – 07/31/2005, \$90,000 total costs.

*NIH Grant CA36622-19-24 "Antineoplastic Agents From Marine Organisms" June 1,
2001 to May 31, 2006, \$1,150,000 (direct costs).*

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American Cancer Society RSG-02-141-01-CNE (David Jones, PI) Retinoid Biosynthesis in Colon Tumorigenesis 07/02/02 – 06/30/06.

ICBG -Fogerty Foundation Conservation and Sustainable Use of Biodiversity of Papua New Guinea (Chris M. Ireland and Louis R. Barrows, co-PIs) 09/01/03 - 8/31/08, \$3,500,000 total costs, 09/25/03 - 05/31/08.

NIH Grant CA67786, 11-15; "Anticancer Agents from Unique Natural Products Sources" September 26, 2005 to April 30, 2010, \$5,920,782 (total costs).

NIH Grant CA36622-24-28 "Antineoplastic Agents From Marine Organisms" December 1, 2006 to November 31, 2011, \$1,250,000 (direct costs).

Publications

1. Gomez, E.; Faulkner, D.J.; Newman, W.A.; Ireland, C. Juvenile Hormone Mimics: Effect on Cirriped Crustacean Metamorphosis *Science* (1973) 179, 813-814.
2. Ramenofsky, M.; Faulkner, D.J.; Ireland, C. Effect of Juvenile Hormone on Cirriped Metamorphosis *Biochem. Biophys. Res. Comm.* (1974) 60, 172-178.
3. Faulkner, D.J.; Stallard, M.O.; Ireland, C.M. Prepacifenol Epoxide, a Halogenated Sesquiterpene Diepoxide *Tetrahedron. Lett.* (1974) 40, 3571-3574.
4. Ireland, C.; Faulkner, D.J.; Clardy, J.; Finer, J. A Novel Diterpene from *Dolabella californica* *J. Amer. Chem. Soc.* (1976) 98, 4664.
5. Ireland, C.M.; Stallard, M.O.; Faulkner, D.J. Some Chemical Constituents of the Digestive Gland of the Sea Hare *Aplysia californica* *J. Org. Chem.* (1976) 41, 2461-2465.
6. Ireland, C. and Faulkner, D.J. Diterpenes from *Dolabella californica* *J. Org. Chem.* (1977) 42, 3157.
7. Ireland, C.; Faulkner, D.J.; Solheim, B.A.; Clardy, J. Tridachione, a Propionate-Derived Metabolite of the Opisthobranch Mollusc *Tridachiella diomedea* *J. Amer. Chem. Soc.* (1978) 100, 1002.
8. Ireland, C. and Faulkner, D.J. The Defensive Secretion of the Opisthobranch Molluscs *Onchidella binneyi* *Bioorg. Chem.* (1978) 7, 125-131.
9. Ireland, C.; Faulkner, D.J.; Clardy, J.; Finer, J. Crispatone, a Metabolite of the Opisthobranch Mollusc *Tridachia crispata* *J. Amer. Chem. Soc.* (1979) 101, 1275.

10. Ireland, C. and Scheuer, P.J. Photosynthetic Marine Molluscs *Science* (1979) 205, 922.
11. Ireland, C. and Scheuer, P.J. Ulicyclamide and Ulithiacylamide, Two New Small Peptides from a Marine Tunicate *J. Amer. Chem. Soc.* (1980) 102, 5688-5691.
12. Ireland, C. and Faulkner, D.J. The Metabolites of the Marine Molluscs *Tridachiella diomedea* and *Tridachia crispata* *Tetrahedron*, (1981) 37, 233-240.
13. Ireland, C.M.; Durso, A.R., Jr.; Scheuer, P.J. N,N¹ dipenethylurea a Metabolite from the Marine Ascidian *Didemnum ternatanum* *J. Nat. Prod.* (1981) 44, 360-361.
14. Hochlowski, J.R.; Walker, R.P.; Ireland, C.M.; Faulkner, D.J. Metabolites of Four Nudibranchs of the Genus *Hypselodoris* *J. Org. Chem.* (1982) 47, 88-91.
15. Ireland, C.M.; Durso, A.R., Jr.; Newman, R.A.; Hacker, M.P. Antineoplastic Cyclic Peptides from the Marine Tunicate *Lissoclinum patella* *J. Org. Chem.* (1982) 47, 1807-1811.
16. Biskupiak, J.E. and Ireland, C.M. Pectinatone, a New Antibiotic from the Mollusc *Siphonaria pectinata* *Tetrahedron Lett.* (1983) 24, 3055-3058.
17. Biskupiak, J.E. and Ireland, C.M. The Stereochemistry of Thiazole Amino Acids in Peptides *J. Org. Chem.* (1983) 48, 2302-2304.
18. Wasylyk, J.M.; Biskupiak, J.E.; Costello, C.E.; Ireland, C.M. Cyclic Peptide Structures from the Tunicate *Lissoclinum patella* by FAB Mass Spectroscopy *J. Org. Chem.* (1983) 48, 4445-4449.
19. Sesin, D.F. and Ireland, C.M. Iodinated Tyramine Derivatives from Marine Tunicate *Tetrahedron Lett.* (1984) 25, 403-404.
20. Ireland, C.M.; Biskupiak, J.E.; Hite, G.J.; Rapposch, M.; Scheuer, P.J.; Ruble, J.R. Ilikonapyrone Esters, Likely Defense Allomones of the Mollusc *Onchidium verruculatum* *J. Org. Chem.* (1984) 49, 559-561.
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23. Roll, D.M. and Ireland, C.M. Citorellamine, A New Bromoindole Derivative from *Polycitorella mariae* *Tetrahedron Lett.* (1985) 26, 4303-4306.

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24. Biskupiak, J.E. and Ireland, C.M. Cytotoxic Metabolites from the Mollusc *Peronia peronii* *Tetrahedron Lett.* (1985) 26, 4307-4310.
25. Sesin, D.F.; Gaskel, S.; Ireland, C.M. The Chemistry of *Lissoclinum patella* *Bull. Soc. Chim. Belg.* (1986) 95, 853-867.
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29. McKee, T.C. and Ireland, C.M. Cytotoxic and Antimicrobial Alkaloids from a Fijian *Xestospongia caycedoi* *J. Nat. Prod.* (1987) 50, 754-756.
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31. Molinski, T.F. and Ireland, C.M. Dysidazirine, a Cytotoxic Azacyclopropene from the Marine Sponge *Dysidea fragilis* *J. Org. Chem.* (1988) 53, 2103.
32. Roll, D.M. and Ireland, C.M. Fascaplysin, an Unusual Antimicrobial Pigment from the Marine Sponge *Fascaplysinopsis* sp. *J. Org. Chem.* (1988) 53, 3276.
33. Ireland, C.M.; Roll, D.M.; Molinski, T.F.; McKee, T.C.; Zabriskie, T.M.; Swersey, J.C. Uniqueness of the Marine Chemical Environment: Categories of Marine Natural Products from Invertebrates *Mem. Cal. Acad. Sci.* (1988) 13, 41-57.
34. Zabriskie, T.M.; Mayne, C.L.; Ireland, C.M. Patellazole C: A Novel Cytotoxic Macrolide from *Lissoclinum patella* *J. Amer. Chem. Soc.* (1988) 110, 7919-7920.
35. McKee, T.C.; Ireland, C.M. The Complete Spectral Assignment of Didemnin B and Nordidemnin B *Tetrahedron Lett.* (1989) 30, 3053-56.
36. Charyulu, G.A.; McKee, T.C.; Ireland, C.M. Diplamine, A Cytotoxic Polyaromatic Alkaloid from the Tunicate *Diplosoma* sp. *Tetrahedron Lett.* (1989) 32, 4201.

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APPENDIX A

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